<u>AMENDMENTS</u>

IN THE CLAIMS:

Please amend the claims as follows:

Please cancel Claims 1-16.

Please add new Claims 17-46 as follows:

- 17. (new) A method for treating Alzheimer's disease in a subject in need of treatment thereof, the method comprising administering to the subject a therapeutic amount of an amidine compound, or a pharmaceutically acceptable salt thereof.
- 18. (new) The method of Claim 17, wherein the amidine comprises a compound of formula (I):

$$A \longrightarrow X \longrightarrow (CH_2)_n \longrightarrow X \longrightarrow B$$

$$R_3 \longrightarrow R_3$$

$$R_3 \longrightarrow R_3$$

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (i):

$$\begin{array}{c}
R_{\uparrow} N \\
R_{\uparrow} N \\
R_{\uparrow} R_{2}
\end{array}$$
(i)

subject to the proviso that at least one of A and B is a compound of formula (i);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkoxyalkyl, cycloalkyl, aryl, hydroxyalkyl, aminoalkyl, and alkylaminoalkyl;

or two R_1 groups on the same compound of formula (i) together represent — $(CH_2)_m$ — wherein m is 2, 3, or 4;

R₃ is H, loweralkyl, oxyalkyl, alkoxyalkyl, hydroxyalkyl, cycloalkyl, aryl, aminoalkyl, alkylaminoalkyl, or halogen;

n is an integer from 2 to 6; and

X is O, NH, or S;

or a pharmaceutically acceptable salt thereof.

19. (new) The method of Claim 18 wherein the amidine comprises a compound selected from the group consisting of:

HN

$$H_2N$$
 $O \leftarrow (CH_2)_{\pi}O$
 NH_2 ;

 NH_2 ;

wherein n is an integer from 2 to 6; or a pharmaceutically acceptable salt thereof.

20. (new) The method of Claim 17, wherein the amidine comprises a compound of formula (II):

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (i):

subject to the proviso that at least one of A and B is a compound of formula (i);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkoxyalkyl, cycloalkyl, aryl, hydroxyalkyl, aminoalkyl, and alkylaminoalkyl;

or two R_1 groups on the same compound of formula (i) together represent — $(CH_2)_m$ — wherein m is 2, 3, or 4;

X is a linear or branched, saturated or unsaturated C_1 - C_{12} alkyl comprising up to 4 double bonds; or X is a heterocyclic aromatic group selected from the group consisting of:

wherein

R₆, R₇, and R₈ are each independently selected from the group consisting of H, loweralkyl, halogen, oxyalkyl, oxyaryl, or oxyarylalkyl;

 R_9 is hydrogen, loweralkyl, hydroxy, aminoalkyl, or alkylaminoalkyl;

or a pharmaceutically acceptable salt thereof.

21. (new) The method of Claim 20, wherein the amidine comprises a compound selected from the group consisting of:

$$HN \longrightarrow NH_2$$
 H_2N
 NH
 H
 N
 N
 N

wherein n is an integer from 1 to 12; or a pharmaceutically acceptable salt thereof.

22. (new) The method of Claim 17, wherein the amidine comprises a compound of formula (III):

$$A \xrightarrow{R_3} (CH_2)_n \xrightarrow{R_3} (III)$$

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (i):

$$\begin{array}{c} R_{1} N \\ \searrow \\ R_{1} N \\ R_{2} \end{array}$$
 (i)

subject to the proviso that at least one of A and B is a compound of formula (i);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkoxyalkyl, cycloalkyl, aryl, hydroxyalkyl, aminoalkyl and alkylaminoalkyl;

or two R_1 groups on the same compound of formula (i) together represent — $(CH_2)_m$ — wherein m is 2, 3, or 4;

or two R_1 groups on the same compound of formula (i) together represent

wherein R₅ is

$$\begin{array}{c}
R_{\uparrow} N \\
\downarrow \\
R_{\uparrow} N \\
\downarrow \\
R_{2}
\end{array}$$
(i)

R₃ is H, loweralkyl, oxyalkyl, alkoxyalkyl, hydroxyalkyl, cycloalkyl, aryl, aminoalkyl, alkylaminoalkyl, or halogen;

n is an integer from 0 to 2; and

 $\,$ X is CH_2O or a heterocyclic aromatic group selected from the group consisting of:

wherein:

R₆, R₇, and R₈ are each independently selected from the group consisting of H, loweralkyl, halogen, oxyalkyl, oxyaryl, or oxyarylalkyl;

R₉ is hydrogen, loweralkyl, hydroxy, aminoalkyl, or alkylaminoalkyl; or a pharmaceutically acceptable salt thereof.

22. (new) The method of Claim 21 wherein the amidine comprises a compound selected from the group consisting of:

or a pharmaceutically acceptable salt thereof.

24. (new) The method of Claim 17, wherein the amidine comprises a compound of formula (IV):

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (i):

subject to the proviso that at least one of A and B is a compound of formula (i);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkoxyalkyl, cycloalkyl, aryl, hydroxyalkyl, aminoalkyl, and alkylaminoalkyl;

or two R_1 groups on the same compound of formula (i) together represent — $(CH_2)_m$ — wherein m is 2, 3, or 4;

or two R_1 groups on the same compound of formula (i) together represent

wherein R₅ is

$$R_{\uparrow} N$$
 (i) $R_{\uparrow} N$ R_{2} ; and

R₃ is H, loweralkyl, oxyalkyl, alkoxyalkyl, hydroxyalkyl, cycloalkyl, aryl, aminoalkyl, alkylaminoalkyl, or halogen; or a pharmaceutically acceptable salt thereof.

25. (new) The method of Claim 24 wherein the amidine comprises a compound selected from the group consisting of:

or a pharmaceutically acceptable salt thereof.

26. (new) The method of Claim 17, wherein the amidine comprises a compound of formula (V):

$$\begin{array}{c|c}
R_4 & B \\
R_3 & R_1 \\
O & R_2
\end{array}$$
(V)

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (ii):

subject to the proviso that at least one of A and B is a compound of formula (ii);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, aryl, alkylaryl, aminoaryl, halogen, oxyalkyl, oxyaryl, or oxyarylalkyl;

 R_3 and R_4 are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkylaryl, aryl, oxyaryl, aminoalkyl, aminoaryl, or halogen;

each R_5 is independently selected from the group consisting of H, loweralkyl, alkoxyalkyl, hydroxyalkyl, aminoalkyl, alkylaminoalkyl, cycloalkyl, aryl, or alkylaryl;

or two R_{5} groups together represent C_{2} to C_{10} alkyl, hydroxyalkyl, or alkylene; and

R₆ is H, hydroxy, loweralkyl, alkoxyalkyl, hydroxyalkyl, aminoalkyl, alkylamino, alkylaminoalkyl, cycloalkyl, hydroxycycloalkyl, alkoxycycloalkyl, aryl, and alkylaryl;

or a pharmaceutically acceptable salt thereof.

27. (new) The method of Claim 17, wherein the amidine comprises a compound of formula (VI):

$$A \longrightarrow (CH_2)_n X \longrightarrow B$$

$$R_3 \longrightarrow R_3$$

$$(VI)$$

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (i):

subject to the proviso that at least one of A and B is a compound of formula (i);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkoxyalkyl, cycloalkyl, aryl, hydroxyalkyl, aminoalkyl, and alkylaminoalkyl;

or two R_1 groups on the same compound of formula (i) together represent — $(CH_2)_m$ — wherein m is 2, 3, or 4;

R₃ is H, loweralkyl, oxyalkyl, alkoxyalkyl, hydroxyalkyl, cycloalkyl, aryl, aminoalkyl, alkylaminoalkyl, or halogen;

or two R_1 groups on the same compound of formula (i) together represent

wherein R₅ is

X is O, S, or NH;

n is an integer from 1 to 8; or a pharmaceutically acceptable salt thereof.

28. (new) The method of Claim 27, wherein the amidine comprises a compound selected from the group consisting of:

$$HN$$
 H_2N
 CH_2
 CH

or a pharmaceutically acceptable salt thereof.

- 29. (new) The method of Claim 17 wherein the amidine comprises a bis-benzamidine.
- 30. (new) The method of Claim 17 wherein the amidine comprises a compound having the following structure:

or a pharmaceutically acceptable salt thereof.

- 31. (new) The method of Claim 17, wherein the subject is afflicted with Alzheimer's disease.
- 32. (new) The method of Claim 17, wherein the subject is at risk of developing Alzheimer's disease, the treatment is a prophylactic treatment, and the amidine compound is administered in a prophylactically effective amount.
- 33. (new) A method for treating diabetes in a subject in need of treatment thereof, the method comprising administering to the subject a therapeutic amount of an amidine compound, or a pharmaceutically acceptable salt thereof.

34. (new) The method of Claim 33, wherein the amidine comprises a compound of formula (I):

$$A \longrightarrow X \longrightarrow (CH_2)_n \longrightarrow X \longrightarrow B$$
 (I)

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (i):

subject to the proviso that at least one of A and B is a compound of formula (i);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkoxyalkyl, cycloalkyl, aryl, hydroxyalkyl, aminoalkyl, and alkylaminoalkyl;

or two R_1 groups on the same compound of formula (i) together represent — $(CH_2)_m$ — wherein m is 2, 3, or 4;

R₃ is H, loweralkyl, oxyalkyl, alkoxyalkyl, hydroxyalkyl, cycloalkyl, aryl, aminoalkyl, alkylaminoalkyl, or halogen;

n is an integer from 2 to 6; and

X is O, NH, or S;

or a pharmaceutically acceptable salt thereof.

35. (new) The method of Claim 34 wherein the amidine comprises a compound selected from the group consisting of:

wherein n is an integer from 2 to 6; or a pharmaceutically acceptable salt thereof.

36. (new) The method of Claim 33, wherein the amidine comprises a compound of formula (II):

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (i):

subject to the proviso that at least one of A and B is a compound of formula (i);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkoxyalkyl, cycloalkyl, aryl, hydroxyalkyl, aminoalkyl, and alkylaminoalkyl;

or two R_1 groups on the same compound of formula (i) together represent — $(CH_2)_m$ — wherein m is 2, 3, or 4;

X is a linear or branched, saturated or unsaturated C_1 - C_{12} alkyl comprising up to 4 double bonds; or X is a heterocyclic aromatic group selected from the group consisting of:

wherein

R₆, R₇, and R₈ are each independently selected from the group consisting of H, loweralkyl, halogen, oxyalkyl, oxyaryl, or oxyarylalkyl;

R₉ is hydrogen, loweralkyl, hydroxy, aminoalkyl, or alkylaminoalkyl; or a pharmaceutically acceptable salt thereof.

37. (new) The method of Claim 36, wherein the amidine comprises a compound selected from the group consisting of:

$$\begin{array}{c} & & & & & & & & \\ & & & & & & & \\ & & & & & & \\ & & & & & & \\ & & & & & & \\ & & & & & \\ & & & & & \\ & & & & & \\ & & & & & \\ & & & & & \\ & & & & \\ & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\ & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & &$$

wherein n is an integer from 1 to 12; or a pharmaceutically acceptable salt thereof.

38. (new) The method of Claim 33, wherein the amidine comprises a compound of formula (III):

$$R_3$$
 $(CH_2)_n$ X $(CH_2)_n$ B (III)

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (i):

subject to the proviso that at least one of A and B is a compound of formula (i);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkoxyalkyl, cycloalkyl, aryl, hydroxyalkyl, aminoalkyl and alkylaminoalkyl;

or two R_1 groups on the same compound of formula (i) together represent — $(CH_2)_m$ — wherein m is 2, 3, or 4;

or two R_1 groups on the same compound of formula (i) together represent

wherein R₅ is

R₃ is H, loweralkyl, oxyalkyl, alkoxyalkyl, hydroxyalkyl, cycloalkyl, aryl, aminoalkyl, alkylaminoalkyl, or halogen;

n is an integer from 0 to 2; and

X is CH₂O or a heterocyclic aromatic group selected from the group consisting of:

wherein:

R₆, R₇, and R₈ are each independently selected from the group consisting of H, loweralkyl, halogen, oxyalkyl, oxyaryl, or oxyarylalkyl;

R₉ is hydrogen, loweralkyl, hydroxy, aminoalkyl, or alkylaminoalkyl; or a pharmaceutically acceptable salt thereof.

39. (new) The method of Claim 38 wherein the amidine comprises a compound selected from the group consisting of:

$$\begin{array}{c} \text{HN} \\ \text{H}_2\text{N} \\ \end{array}, \\ \text{HN} \\ \text{S} \\ \end{array}, \\ \text{NH}_2 \\ \vdots \\ \text{NH}_2 \\ \vdots \\ \end{array}$$

$$\begin{array}{c} HN \\ H_{2}N \end{array}$$

$$\begin{array}{c} NH \\ NH_{2} \end{array}$$

$$\begin{array}{c} NH \\ NH_{2} \end{array}$$

$$\begin{array}{c} NH \\ NH_{2} \end{array}$$

$$\begin{array}{c} CH_{3} \\ NH \end{array}$$

$$\begin{array}{c} CH_{3} \\ CH_{3} \end{array}$$

$$\begin{array}{c} CH_{3} \\ NH \end{array}$$

$$\begin{array}{c} CH_{3} \\ NH \end{array}$$

$$\begin{array}{c} HN \\ NH \end{array}$$

$$\begin{array}{c} NH \\ NH \end{array}$$

or a pharmaceutically acceptable salt thereof.

40. (new) The method of Claim 33, wherein the amidine comprises a compound of formula (IV):

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (i):

$$\begin{array}{ccc}
R_{7} & N \\
R_{7} & N \\
R_{2} & \end{array}$$

subject to the proviso that at least one of A and B is a compound of formula (i);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkoxyalkyl, cycloalkyl, aryl, hydroxyalkyl, aminoalkyl, and alkylaminoalkyl;

or two R_1 groups on the same compound of formula (i) together represent — $(CH_2)_m$ — wherein m is 2, 3, or 4;

or two R_1 groups on the same compound of formula (i) together represent

wherein R5 is

$$R_{\uparrow} N$$
 (i) $R_{\uparrow} N$ R_{2} ; and

R₃ is H, loweralkyl, oxyalkyl, alkoxyalkyl, hydroxyalkyl, cycloalkyl, aryl, aminoalkyl, alkylaminoalkyl, or halogen; or a pharmaceutically acceptable salt thereof.

40. (new) The method of Claim 39 wherein the amidine comprises a compound selected from the group consisting of:

or a pharmaceutically acceptable salt thereof.

41. (new) The method of Claim 33, wherein the amidine comprises a compound of formula (V):

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (ii):

$$\begin{array}{c}
R_{5} N \\
R_{5} N \\
R_{6}
\end{array}$$
(ii)

subject to the proviso that at least one of A and B is a compound of formula (ii);

 R_1 and R_2 are each independently selected from the group consisting of H, loweralkyl, aryl, alkylaryl, aminoaryl, halogen, oxyalkyl, oxyaryl, or oxyarylalkyl;

R₃ and R₄ are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkylaryl, aryl, oxyaryl, aminoalkyl, aminoaryl, or halogen;

each R_5 is independently selected from the group consisting of H, loweralkyl, alkoxyalkyl, hydroxyalkyl, aminoalkyl, alkylaminoalkyl, cycloalkyl, aryl, or alkylaryl;

or two R_5 groups together represent C_2 to C_{10} alkyl, hydroxyalkyl, or alkylene; and

R₆ is H, hydroxy, loweralkyl, alkoxyalkyl, hydroxyalkyl, aminoalkyl, alkylamino, alkylaminoalkyl, cycloalkyl, hydroxycycloalkyl, alkoxycycloalkyl, aryl, and alkylaryl;

or a pharmaceutically acceptable salt thereof.

42. (new) The method of Claim 33, wherein the amidine comprises a compound of formula (VI):

$$A \longrightarrow (CH_2)_n X \longrightarrow B$$

$$R_3 \longrightarrow R_3$$
(VI)

wherein:

A and B are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, nitro, amino, aminoalkyl, halo, hydroxy, carboxy, and compounds of formula (i):

$$\begin{array}{c} R_{\uparrow} N \\ \downarrow \\ R_{\uparrow} N \\ \downarrow \\ R_{2} \end{array}$$
 (i)

subject to the proviso that at least one of A and B is a compound of formula (i);

R₁ and R₂ are each independently selected from the group consisting of H, loweralkyl, oxyalkyl, alkoxyalkyl, cycloalkyl, aryl, hydroxyalkyl, aminoalkyl, and alkylaminoalkyl;

or two R_1 groups on the same compound of formula (i) together represent — $(CH_2)_m$ — wherein m is 2, 3, or 4;

R₃ is H, loweralkyl, oxyalkyl, alkoxyalkyl, hydroxyalkyl, cycloalkyl, aryl, aminoalkyl, alkylaminoalkyl, or halogen;

or two R_1 groups on the same compound of formula (i) together represent

wherein R₅ is

$$\begin{array}{c} R_{\uparrow} N \\ \searrow & \text{(i)} \\ R_{\uparrow} N \\ R_{z} \end{array}$$

X is O, S, or NH;

n is an integer from 1 to 8; or a pharmaceutically acceptable salt thereof.

44. (new) The method of Claim 43, wherein the amidine comprises a compound selected from the group consisting of:

$$\begin{array}{c} \text{HN} \\ \text{H}_2 \text{N} \end{array} \longrightarrow \begin{array}{c} \text{CH}_2 - \text{O} \\ \end{array} \longrightarrow \begin{array}{c} \text{NH} \\ \text{NH}_2 \\ \vdots \end{array}$$

$$H_2N$$
 CH_2-O

$$\begin{array}{c} \text{NH} \\ \text{NH}_2; \\ \text{HN} \\ \text{H}_2 \text{N} \\ \end{array}; \\ \text{CH}_2 - \text{O} \\ \text{CH}_2 - \text{O} \\ \text{NH}_2; \\ \text{SIMP} \\ \text{SIMP}$$

or a pharmaceutically acceptable salt thereof.

- 45. (new) The method of Claim 33 wherein the amidine comprises a bis-benzamidine.
- 46. (new) The method of Claim 33 wherein the amidine comprises a compound having the following structure:

or a pharmaceutically acceptable salt thereof.

- 47. (new) The method of Claim 33, wherein the subject is afflicted with diabetes.
- 48. (new) The method of Claim 33, wherein the subject is at risk of developing diabetes, the treatment is a prophylactic treatment, and the amidine compound is administered in a prophylactically effective amount.